## PATENT COOPERATION TREATY

# **PCT**

### INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's tile reference	FOR FURTHER		see Form PCT/ISA/220
SCB/PB60728	ACTION		il as, where applicable, item 5 below.
nternational application No.	International filing date (day/mor	lh/year)	(Earliest) Priority Date (day/month/year)
CT/GB2005/000348	01/02/2009	5	03/02/2004
pplicant	· · · · · · · · · · · · · · · · · · ·		
7.340 CD CD CD C			
LAXO GROUP LIMITED	<u> </u>		<u> </u>
This international Search Report has be according to Article 18. A copy is being t	en prepared by this international Ses ransmitted to the international Bures	erching Aut U.	hority and is transmitted to the applicant
This International Search Report consist	s of a total ofsh	eels.	
	y a copy of each prior art document		report.
1 Boole of the report	<u> </u>	-	<u> </u>
<ol> <li>Basis of the report</li> <li>a. With regard to the language, the</li> </ol>	a International search was carried ou	on the bas	sis of the international application in the
language in which it was filed, ur	ness otherwise indicated under this i	tem.	ere of the international application in the
The international this Authority (Re	l search was carried out on the basis ule 23.1(b)).	of a transli	ation of the international application furnished to
b. With regard to any nucle	otide and/or amino acid sequence	disclosed	in the international application, see Box No. 1.
	ınd unsearchable (See Box II).		
	ma ansasichanta (266 Box II).		
Unity of invention is lac	king (see Box III).		
. With regard to the tille,			
the lext is approved as su	ubmitted by the applicant.		
	shed by this Authority to read as follo		
COMBINATION OF CB2 MOI	DULATORS AND PDE4 INHI	BITORS	FOR USE IN MEDICINE
With regard to the abstract,			
the text is approved as su	bmitted hv the ≏notices•		
X the text has been establish	hed according to Bule 38 9/6). No 46	ls Authorite	as It appears in Box No. IV. The applicant
may, within one month tro	m the date of mailing of this internal	onal search	r as it appears in Box No. IV. The applicant I report, submit comments to this Authority.
With regard to the drawings,			
a. the floure of the drawings to be o	ublished with the abstract is Figure N	lo,	
Para an an an an an an an b	The second of the land of the		<del></del>
as suggested by t			
as suggested by t	he applicant,	led to sugg	est a ligure.
as suggested by this			<del>-</del>

International application No.

PCT/GB2005/000348

#### Box No. IV Text of the abstract (Continuation of item 5 of the first sheet)

Combination of one or more CB2 modulators such as a compound of formula (I), (II) and (III)

and one or more PDE4 inhibitors are useful of treating conditions which are mediated by the activity of CB2 receptors or conditions which are mediated by PDE4, such as an immune disorder, an inflammatory disorder, pain, rheumatoid

Form PCT/ISA/210 (continuation of first sheet (3)) (January 2004)

JUL. 28. 2006 10:26AM GW GLOBAL R&D IS

International Application No PCT/GB2005/000348

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 A61K31/505 A61P11/00 A61P37/00 A61P29/00

According to International Patent Classification (IPC) or to both national classification and IPC

#### B. FIELDS SEARCHED

Minimum documentation sparched (classification system followed by classification symbols) IPC 7 A61K

Documentation searched other than minimum documentation to the extent that such documents are included. In the fields searched

Electronic data base consulted during the international search (name of data base and, where precilcal, search terms used)

EPO-Internal, WPI Data, EMBASE, BIOSIS, PAJ

Catégory *	Criation of document, with indication, where appropriate, of the relovant passages	Relevant to claim No.
Y	FR 2 839 718 A (SANOFI-SYNTHELABO) 21 November 2003 (2003-11-21) abstract page 13, line 10 - page 14, line 5	1-7
Y	WO 02/060447 A (UNIVERSITY OF CONNECTICUT; MAKRIYANNIS, ALEXANDROS; DENG, HONGFENG) 8 August 2002 (2002-08-08) page 8, line 7 - line 18 page 2, line 1 - line 8	1-7
Y	US 5 925 768 A (BARTH ET AL) 20 July 1999 (1999-07-20) column 1, paragraph 2 column 26, line 11 - line 29	1-7
	-/	,

X Further documents are listed in the continuation of box C.	Patent (amily members are listed in annex.
<ul> <li>Special categories of cited documents:</li> <li>"A" document defining the general state of the last which is not considered to be of particular relevance</li> <li>"E" earlier document but published on or after the International filling date</li> <li>"L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or either special reason (as specified)</li> <li>"O" document referring to an oral disclosure, use, exhibition or other means</li> <li>"P" document published prior to the international filling date but faler than the priority date claimed</li> </ul>	<ul> <li>'T' later document published after the international flling dete or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention</li> <li>"X' document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone</li> <li>"Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such document is combined with one or more other such documents, such combination being obvious to a person skilled in the art.</li> <li>"&amp;" document member of the same patent family</li> </ul>
Date of the actual completion of the international search  22 April 2005	Date of mailing of the informational search report  06/05/2005
Name and malling address of the ISA  European Palent Office, P.B. 5815 Patentlaan 2  NL = 2260 HV Rijswijk  Tel. (+31=70) 340-2040, Tx. 31 651 apo nt,  Fax: (+31-70) 340-3016	Authorized officer Glacobbe, S

International Application No PCT/GB2005/000348

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Délouge de dels Ma
		Rélevant to claim No.
Υ	WO 97/09315 A (SIGNAL PHARMACEUTICALS, INC; SUTO, MARK, J; GOLDMAN, MARK, E; GAYO, LE) 13 March 1997 (1997-03-13) claims 1,36-44	4,5
Υ	HUANG Z ET AL: "The next generation of PDE4 inhibitors." CURRENT OPINION IN CHEMICAL BIOLOGY. AUG 2001, vol. 5, no. 4, August 2001 (2001-08), pages 432-438, XP002325704 ISSN: 1367-5931 figure 1 page 432, column 1, paragraph 1	1-7
Y	SOUNESS J E ET AL: "Immunosuppressive and anti-inflammatory effects of cyclic AMP phosphodiesterase (PDE) type 4 inhibitors."  IMMUNOPHARMACOLOGY. MAY 2000, vol. 47, no. 2-3, May 2000 (2000-05), pages 127-162, XP002325705  ISSN: 0162-3109 the whole document figures 3,4	1-7
Y	WO 02/062750 A (SCHERING CORPORATION) 15 August 2002 (2002-08-15) page 39, line 13	1-7
<b>f</b>	WO 2004/000807 A (SCHERING CORPORATION; KOZLOWSKI, JOSEPH, A; SHANKAR, BANDARPALLE, B; S) 31 December 2003 (2003-12-31) page 29, line 27	17
Y , Y	WO 2004/018433 A (GLAXO GROUP LIMITED; EATHERTON, ANDREW; JOHN; GIBLIN, GERARD, MARTIN,) 4 March 2004 (2004-03-04) the whole document	1-7
, Y	WO 2004/018434 A (GLAXO GROUP LIMITED; EATHERTON, ANDREW, JOHN; GIBLIN, GERARD, MARTIN,) 4 March 2004 (2004-03-04) the whole document	1-7
, Y	WO 2004/029026 A (GLAXO GROUP LIMITED; EATHERTON, ANDREW, JOHN; GIBLIN, GERARD, MARTIN,) 8 April 2004 (2004-04-08) the whole document	1-7
, Y	WO 2004/029027 A (GLAXO GROUP LIMITED; EATHERTON, ANDREW, JOHN; GIBLIN, GERARD, MARTIN,) 8 April 2004 (2004-04-08) the whole document	1-7
1	<b></b>	

# GW GLOBAL R&D IS INTERNATIONAL SEARCH REPORT

international Application No PCT/GB2005/000348

Citation of document, with Indication, where appropriate, of the relevant passages	Relevant to claim No.
WO 2004/085385 A (SCHERING CORPORATION; SHANKAR, BANDARPALLE, B; RIZVI, RAZIA, K; KOZLOW) 7 October 2004 (2004-10-07) page 28, line 19 - line 25	1-3,6,7
	,
	WO 2004/085385 A (SCHERING CORPORATION; SHANKAR, BANDARPALLE, B; RIZVI, RAZIA, K; KOZLOW) 7 October 2004 (2004-10-07)

International application No. PCT/GB2005/000348

Box II Observations where certain claims were found unsearchable (Continuation of Item 2 of first sheet)	
This International Search Report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons;	
1. X Claims Nos.: because they relate to subject matter not required to be searched by this Authority, namely:	
Although claims 1, 2 and 4-6 are directed to a method of treatment of the human/animal body, the search has been carried out and based on the alleged effects of the compound/composition.	
2. Claims Nos.; because they relate to parts of the International Application that do not comply with the prescribed requirements to such an extent that no meaningful International Search can be carried out, specifically:	
3. Claims Nos.; because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).	
Box III Observations where unity of invention is lacking (Continuation of Item 3 of first sheet)	
This International Searching Authority found multiple Inventions in this International application, as follows: .	
1. As all required additional search fees were timely paid by the applicant, this international Search Report covers all searchable claims.	
2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment of any additional fee.	
3. As only some of the required additional search fees were timely paid by the applicant, this international Search Report covers only those claims for which fees were paid, specifically claims Nos.:	
4. No required additional search fees were timely paid by the applicant. Consequently, this international Search Report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:	
Remark on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.	

Information on petent family members

PCT/GB2005/000348

	<del></del>			PC1/GB2005/000348			
Patent document cited in search report		Publication date		Patent family member(s)	Publication date		
FR 2839718	A	21-11-2003	FR AU BR CA EP WO HR	2839718 A1 2003258762 A1 0311185 A 2482756 A1 1507758 A2 03097597 A2 20041073 A2	21-11-2003 02-12-2003 22-02-2005 27-11-2003 23-02-2005 27-11-2003 31-12-2004		
WO 02060447	Α	08-08-2002	CA EP JP WQ	2436133 A1 1363632 A1 2005503998 T 02060447 A1	08-08-2002 26-11-2003 10-02-2005 08-08-2002		
U\$ 5925768	A	20-07-1999	FATUURANZEEKEEWGHJJNNRPRUSKRWA	2742148 A1 191211 T 718763 B2 1101097 A 9611986 A 2239489 A1 1207731 A ,C 9801775 A3 69607484 D1 69607484 T2 868420 T3 9800171 A 0868420 A1 0885889 A2 2148820 T3 9721682 A1 3033626 T3 9721682 A1 3033626 T3 9901244 A2 2000502080 T 3510270 B2 982589 A 323962 A 327177 A1 868420 T 2170230 C2 868420 T1 75898 A3 9801041 T2 402594 B 9610299 A	13-06-1997 15-04-2000 20-04-2000 03-07-1997 17-02-1999 19-06-1997 10-02-1999 14-10-1998 04-05-2000 26-10-2000 21-08-2000 15-12-1998 07-10-1998 23-12-1998 16-10-2000 19-06-1997 31-10-2000 30-08-1999 22-02-2000 22-03-2004 03-08-1998 28-05-1999 23-11-1998 31-08-2000 10-07-2001 31-10-2000 11-01-1999 21-08-1998 21-08-2000 18-06-1997		
WO 9709315	A	13-03-1997	US AU AU AU CA EP JP WO WO US	5811428 A 726058 B2 7013096 A 726522 B2 7163196 A 2230894 A1 2230896 A1 0850228 A1 11512390 T 11512399 T 9709325 A1 9709315 A1 5935966 A	22-09-1998 26-10-2000 27-03-1997 09-11-2000 27-03-1997 13-03-1997 13-03-1997 01-07-1998 26-10-1999 26-10-1999 13-03-1997 13-03-1997 13-03-1997		

information on patent family members

International Application No PCT/GB2005/000348

				1 017 0020037 00 0340				
	tent document in search report		Publication date		Patent (amily member(s)		Publication date	
WO	02062750	A	15-08-2002	BR	0206955	Δ	09-03-2004	
				ÇA	2436659		15-08-2002	
				ČN	1489576		14-04-2004	
				čZ	20032122		15-10-2003	
				ĔΡ	1368308	•		
				ΗŪ	0304060		10-12-2003	
				JP	2004530649	T	28-04-2004	
				ΜX	PA03007078	n n	07-10-2004	
				NO	20033505		18-11-2003	
				SK	9952003		07-10-2003	
				WO			08-01-2004	
				U\$	02062750		15-08-2002	
				UŞ	2003096844		22-05-2003	
		<b></b>			2003232859	AI	18-12-2003	
WO 2	2004000807	A	31-12-2003	ΑU	2003243637	 A1	06-01-2004	
				CA	2487346		31-12-2003	
				WO	2004000807		31-12-2003	
				US	2004044051		04-03-2004	
WO 2	2004018433	 А	04-03-2004	 AU	2002264076	 		
		24	VT VJ ZVU4	MO	2003264076		11-03-2004	
		<b>-</b>		WV	2004018433	A1	04-03-2004	
WO 2	2004018434	Α	04-03-2004	AU	2003260436	<u>-</u> Δ1	11-03-2004	
				WO	2004018434		04-03-2004	
						'		
WO 2	004029026	Α	08 <b>-</b> 0 <b>4</b> -2 <b>00</b> 4	ΑU	2003268907 /	41	19-04-2004	
				WO	2004029026 /		08-04-2004	
WU 2	004029027	Α	08- <b>04-</b> 2004	ΑU	2003276028 #	۱۱	19-04-2004	
<b>-</b>				MO	2004029027	11	08-04-2004	
WO 2	004085385	A	07-10-2004		2004206340	- <b>-</b>	**************************************	
<b>-</b>		Л	V/ -10-2004	US	2004186148 #	- —	23-09-2004	
				WO	2004085385 A	1Z	07-10-2004	